#### PCT -

# 国際事務局 特許協力条約に基づいて公開された国際出願

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(51) 国際特許分類6 WO98/37757 (11) 国際公開番号 A1 A01K 67/027 1998年9月3日(03.09.98) (43) 国際公開日 (21) 国際出願番号 PCT/JP98/00860 (74) 代理人 弁理士 平木祐輔, 外(HIRAKI, Yusuke et al.) 1998年3月2日(02.03.98) (22) 国際出願日 〒105-0001 東京都港区虎ノ門一丁目17番1号 虎ノ門5森ビル3F Tokyo, (JP) (30) 優先権データ 特願平9/62309 1997年2月28日(28.02.97) AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, (81) 指定国 CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, (71) 出願人(米国を除くすべての指定国について) ID, IL, IS, JP, KE, KG, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, 麒麟麦酒株式会社(KIRIN BEER KABUSHIKI KAISHA)[JP/JP] SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW, 〒104-8288 東京都中央区新川二丁目10番1号 Tokyo, (JP) ARIPO特許 (GH, GM, KE, LS, MW, SD, SZ, UG, ZW), ユーラシ (72) 発明者;および ア特許 (AM, AZ, BY, KG, KZ, MD, RU, TJ, TM), 欧州特許 (75) 発明者/出願人(米国についてのみ) (AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, 富塚一磨(TOMIZUKA, Kazuma)[JP/JP] PT, SE), OAPI特許 (BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, 吉田 均(YOSHIDA, Hitoshi)[JP/JP] NE, SN, TD, TG). 石田 功(ISHIDA, Isao)[JP/JP] 〒236-0004 神奈川県横浜市金沢区福浦1-13-5 添付公開書類

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国際調査報告書

(54) Title: PLURIPOTENT CELLS HAVING DISRUPTED INTRINSIC GENES

(54)発明の名称 / 内因性遺伝子が破壊されている分化多能性保持細胞

麒麟麦酒株式会社 基盤技術研究所内 Kanagawa, (JP)

花岡和則(HANAOKA, Kazunori)[JP/JP] 〒228-8555 神奈川県相模原市北里1-15-1 北里大学 理学部 生物科学科内 Kanagawa, (JP)

押村光雄(OSHIMURA, Mitsuo)[JP/JP] 〒683-8503 鳥取県米子市西町86

鳥取大学 医学部 生命科学科内 Tottori, (JP)

#### (57) Abstract

A process for constructing a chimeric non-human animal characterized by preparing microcells containing one or more foreign chromosomes or fragments thereof, and transferring the one or more foreign chromosomes or fragments thereof into pluripotent cells through fusion with the microcells; chimeric non-human animals which can be constructed by the above process and offspring thereof; tissues and cells originating in the above animals; methods for using the individuals, tissues and cells of the animals; pluripotent cells containing one or more foreign chromosomes or fragments thereof, a process for constructing these cells, and the use of these cells; pluripotent cells in which at least two intrinsic genes have been disrupted; a process for constructing the above cells by homologous recombination; etc.

# INTERNATIONAL SEARCH REPORT

International application No.
PCT/JP98/00860

A. CLASSIFICATION OF SUBJECT MATTER Int.Cl <sup>6</sup> A01K67/027					
According to	International Patent Classification (IPC) or to both nat	tional classification and IPC			
B. FIELDS	SEARCHED		· .		
Minimum documentation searched (classification system followed by classification symbols) Int.Cl <sup>6</sup> A01K67/027					
Documentati	Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched				
Electronic data base consulted during the international search (name of data base and, where practicable, search terms used) JICST File on Science and Technology BIOSIS PREVIEWS					
C. DOCUI	MENTS CONSIDERED TO BE RELEVANT				
Category*	Citation of document, with indication, where app		Relevant to claim No.		
Р, Х	Nature Genetics, 16[2] (1997) Full text	) p.133-143	1-40, 42, 43, 45-53		
P', Y	Full text	·	41, 44, 54-76		
Y	JP, 05-328878, A (The Instit Chemical Research), December 14, 1993 (14. 12. 93 Full text (Family: none)	,	1-76		
Y	JP, 07-067629, A (K.K. Dinad March 14, 1995 (14. 03. 95), Full text (Family: none)	io),	1-76		
Y	EMBO Journal, 2 (1983) p.1963 Full text	3-1968	1-76		
Y	Proc. Natl. Acad. Sci. USA, Full text	74 (1997) p.319-323	1-76		
× Furthe	Further documents are listed in the continuation of Box C. See patent family annex.				
"A" document defining the general state of the art which is not considered to be of particular relevance "E" earlier document but published on or after the international filing date "L" document which may throw doubts on priority claim(s) or which is cited to establish the publication date of another citation or other special reason (as specified) "O" document referring to an oral disclosure, use, exhibition or other means "P" document published prior to the international filing date but later than the priority date claimed		"T" later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention  "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone  "Y" document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art document member of the same patent family			
Date of the actual completion of the international search May 21, 1998 (21. 05. 98)  Date of mailing of the international search June 2, 1998 (02. 06. 98)					
Name and mailing address of the ISA/ Japanese Patent Office		Authorized officer			
Facsimile No.		Telephone No.			

# INTERNATIONAL SEARCH REPORT

International application No.
PCT/JP98/00860

ategory*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
Y A	WO, 89/05308, A1 (GESELLSCHAFT FUR BIOTECHNOLOGISCHE FORSCHUNG MBH), June 15, 1989 (15. 06. 89), Full text Full text & JP, 02-502426, A	41, 44, 54, 55 1-40, 42, 43, 45-53, 56-63
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発明の属する分野の分類(国際特許分類(IPC)) Int. Cl A01K67/027 B. 調査を行った分野 調査を行った最小限資料(国際特許分類(IPC)) Int. Cl A01K67/027 最小限資料以外の資料で調査を行った分野に含まれるもの 国際調査で使用した電子データベース(データベースの名称、調査に使用した用語) 科学技術文献ファイル JICST BIOSIS PREVIEWS 関連すると認められる文献 関連する 引用文献の 請求の範囲の番号 引用文献名 及び一部の箇所が関連するときは、その関連する箇所の表示 カテゴリー\* Nature Genetics, 16[2](1997) p. 133-143 1-40, 42, 43, P, X 全文 45-53 41, 44, 54-76 全文 P, Y JP,05-328878,A(理化学研究所) 1-76 Y 14. 12月. 1993 (14. 12. 93) 全文(ファミリーなし) | C欄の続きにも文献が列挙されている。 \* 引用文献のカテゴリー の日の後に公表された文献 「T」国際出願日又は優先日後に公表された文献であって 「A」特に関連のある文献ではなく、一般的技術水準を示す て出願と矛盾するものではなく、発明の原理又は理 もの 論の理解のために引用するもの 「E」先行文献ではあるが、国際出願日以後に公表されたも 「X」特に関連のある文献であって、当該文献のみで発明 「L」優先権主張に疑義を提起する文献又は他の文献の発行 の新規性又は進歩性がないと考えられるもの 「Y」特に関連のある文献であって、当該文献と他の1以 日若しくは他の特別な理由を確立するために引用する 上の文献との、当業者にとって自明である組合せに 文献 (理由を付す) よって進歩性がないと考えられるもの 「O」ロ頭による開示、使用、展示等に含及する文献 「&」同一パテントファミリー文献 「P」国際出願日前で、かつ優先権の主張の基礎となる出願 国際調査報告の発送日 02.06.98 国際調査を完了した日 21.05.98 9414 特許庁審査官(権限のある職員)・・・・・ 国際調査機関の名称及びあて先 哪們 日本国特許庁(ISA/JP) 関根 裕 郵便番号100-8915 東京都千代田区霞が関三丁目4番3号 電話番号 03-3581-1101 内線 3238

C (続き). 関連すると認められる文献				
引用文献の カテゴリー*	引用文献名 及び一部の箇所が関連するときは、その関連する箇所の表示	関連する 請求の範囲の番号		
Y	JP, 07-067629, A (株式会社ディナード) 14.3月.1995 (14.03.95) 全文 (ファミリーなし)	1-76		
Y	EMBO Journal, 2 (1983) p. 1963-1968 全文	1-76		
Y	Proc. Natl. Acad. Sci. USA, 74 (1977) p. 319-323 全文	1-76		
Y A	WO, 89/05308, A1 (GESELLSCHAFT FUR BIOTECHNOLOGISCHE FORSCHUNG MBH) 15.6月.1989 (15.06.89) 全文 全文 全文	41, 44, 54, 55 1-40, 42, 43, 45-53, 56-63		
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#### Derwent abstract of WO 98/37757

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             **Image available**
WPI Acc No: 1998-480821/199841
XRAM Acc No: C98-145439
XRPX Acc No: N98-375183
 Pluripotent cells containing foreign chromosomes or fragments - and
non-human chimeric animals constructed using them and expressing foreign
 genes such as human antibiotic genes
Patent Assignee: KIRIN BEER KK (KIRI )
Inventor: HANAOKA K; ISHIDA I; OSHIMURA M; TOMIZUKA K; YOSHIDA H
Number of Countries: 081 Number of Patents: 004
Patent Family:
                             Applicat No
                                            Kind
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Patent No
              Kind
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              A1 19980903 WO 98JP860
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WO 9837757
                   19980918 AU 9861189
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EP 972445
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                             WO 98JP860
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Priority Applications (No Type Date): JP 9762309 A 19970228
Patent Details:
Patent No Kind Lan Pq
                         Main IPC
                                     Filing Notes
WO 9837757
             A1 J 217 A01K-067/027
   Designated States (National): AL AM AT AU AZ BA BB BG BR BY CA CH CN CU
   CZ DE DK EE ES FI GB GE GH GM GW HU ID IL IS JP KE KG/ KR KZ LC LK LR LS
   LT LU LV MD MG MK MN MW MX NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR
   TT UA UG US UZ VN YU ZW
   Designated States (Regional): AT BE CH DE DK EA ES FI FR GB GH GM GR IE
   IT KE LS LU MC MW NL OA PT SD SE SZ UG ZW
AU 9861189
                       A01K-067/027
                                     Based on patent WO 9837757
              Α
EP 972445
                       A01K-067/027 Based on patent WO 9837757
              A1 E
   Designated States (Regional): AT BE CH DE DK ES FI FR GB GR IE IT LI LU
   MC NL PT SE
                       A01K-067/027 Based on patent WO 9837757
JP 10537525
              Х
Abstract (Basic): WO 9837757 A
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Pluripotent cells containing foreign chromosomes or their fragments (preferably at least 670 kb in length, especially more than 1000 kb) are obtained by preparing cancerous cells containing the foreign chromosomes or fragments, then fusing these with pluripotent cells such as embryonic stem cells, embryonic reproductive cells, embryonic cancer cells or their mutants.

Also claimed are: (1) a method of obtaining hybridoma cells by fusing a cell with a high ability to produce hybridoma cells (such as mouse A9 cells) with a cell containing the foreign chromosomes or fragments (such as normal human diploid cells); (2) a method of utilising pluripotent cells to produce chimeric and transgenic non-human animals (especially mammals such as mice) which can express

the foreign chromosomes or fragments introduced; and (3)chimeric animals, their offspring and tissues and cells derived from the offspring produced by a method as in (2).

USE - The inventions can be used for the production of monoclonal antibodies for medical use which are of human type and therefore not antigenic in humans. They can also be used in the production of chimeric and transgenic animals which express useful foreign proteins, or which can serve as models for the study of human diseases.

Dwg.10/45

Title Terms: CELL; CONTAIN; FOREIGN; CHROMOSOME; FRAGMENT; NON; HUMAN; CHIMERIC; ANIMAL; CONSTRUCTION; EXPRESS; FOREIGN; GENE; HUMAN; ANTIBIOTIC : GENE

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Derwent Class: B04; D16; P14

International Patent Class (Main): A01K-067/027

File Segment: CPI; EngPI

Manual Codes (CPI/A-N): B04-F01; B04-G21; D05-H11A1; D05-H15; D05-H16A

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Chemical Fragment Codes (M1):

\*01\* M423 M710 M903 N135 Q233 V600 V645 V753 V754

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